SEQUENCE LISTING

Oppermann, Hermann Tai, Mei-Sheng McCartney, John

<120> Modified TGF-beta Superfamily Proteins

<130> STK-075

<140> US 09/375,333

<141> 1999-08-16

<160> 124

<170> PatentIn version 2.0

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Gly Cys His
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Arg Ser Leu Arg Ala Ala Ala Pro His Ser Phe Val Ala Leu Trp Ala
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ccc ctg ttc ctg ctg cgc tcc gcc ctg gcc gac ttc agc ctg gac aac
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Pro Leu Phe Leu Leu Arg Ser Ala Leu Ala Asp Phe Ser Leu Asp Asn
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			gtc Val							489
			cat His	-	-					537
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			ttc Phe 185							633
			cac His							681
			tgg Trp							729
			aac Asn							777
			gtg Val							825
			att Ile 265							873

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														agc Ser		1017
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	_				_			_	_	_				atc Ile		1209
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			atg Met									tago	etect	ccc		1351
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Leu Asp Asn 35	. Glu Val His	Ser Ser Phe	Ile His Arg	Arg Leu Arg Ser 45										
Gln Glu Arg 50	Arg Glu Met	Gln Arg Glu 55	Ile Leu Ser 60	Ile Leu Gly Leu										
Pro His Arg 65	Pro Arg Pro	His Leu Gln	Gly Lys His 75	Asn Ser Ala Pro 80										
Met Phe Met	Leu Asp Leu 85	Tyr Asn Ala	Met Ala Val 90	Glu Glu Gly Gly 95										
Gly Pro Gly	Gly Gln Gly 100	Phe Ser Tyr	Pro Tyr Lys	Ala Val Phe Ser 110										
Thr Gln Gly 115		Ala Ser Leu 120	Gln Asp Ser	His Phe Leu Thr 125										
Asp Ala Asp 130	Met Val Met	Ser Phe Val	Asn Leu Val	Glu His Asp Lys										
Glu Phe Phe 145	His Pro Arg 150	Tyr His His	Arg Glu Phe 155	Arg Phe Asp Leu 160										
Ser Lys Ile	Pro Glu Gly 165	Glu Ala Val	Thr Ala Ala 170	Glu Phe Arg Ile 175										
Tyr Lys Asp	Tyr Ile Arg 180	Glu Arg Phe 185	Asp Asn Glu	Thr Phe Arg Ile 190										
Ser Val Tyr 195		Gln Glu His 200	Leu Gly Arg	Glu Ser Asp Leu 205										
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215

220

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His	Asn	Leu	Gly	Leu 245	Gln	Leu	Ser	Val	Glu 250	Thr	Leu	Asp	Gly	Gln 255	Ser
Ile	Asn	Pro	Lys 260	Leu	Ala	Gly	Leu	Ile 265	Gly	Arg	His	Gly	Pro 270	Gln	Asn
Lys	Gln	Pro 275	Phe	Met	Val	Ala	Phe 280	Phe	Lys	Ala	Thr	Glu 285	Val	His	Phe
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Gly	Tyr	Ala 355	Ala	Tyr	Tyr	Cys	Glu 360	Gly	Glu	Сув	Ala	Phe 365	Pro	Leu	Asn
Ser	Tyr 370	Met	Asn	Ala	Thr	Asn 375	His	Ala	Ile	Val	Gln 380	Thr	Leu	Val	His
Phe 385	Ile	Asn	Pro	Glu	Thr 390	Val	Pro	Lys	Pro	Суs 395	Сув	Ala	Pro	Thr	Gln 400
Leu	Asn	Ala	Ile	Ser 405	Val	Leu	Tyr	Phe	Asp 410	Asp	Ser	Ser	Asn	Val 415	Ile

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Lys Trp Ile His Glu Pro Lys Gly Tyr His Ala Asn Phe Cys Leu Gly 25

Pro Cys Pro Tyr Ile Trp Ser Leu Asp Thr Gln Tyr Ser Lys Val Leu

Ala Leu Tyr Asn Gln His Asn Pro Gly Ala Ser Ala Ala Pro Cys Cys

Val Pro Gln Ala Leu Glu Pro Leu Pro Ile Val Tyr Tyr Val Gly Arg

Lys Pro Lys Val Glu Gln Leu Ser Asn Met Ile Val Arg Ser Cys Lys

Cys Ser

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Ala Cys Pro Tyr Leu Trp Ser Ser Asp Thr Gln His Ser Arg Val Leu

Ser Leu Tyr Asn Thr Ile Asn Pro Glu Ala Ser Ala Ser Pro Cys Cys 50 55 60

Val Ser Gln Asp Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Ile Gly Lys

Thr Pro Lys Ile Glu Gln Leu Ser Asn Met Ile Val Lys Ser Cys Lys 90 85

Cys Ser

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Lys Trp Val His Glu Pro Lys Gly Tyr Tyr Ala Asn Phe Cys Ser Gly

Pro Cys Pro Tyr Leu Arg Ser Ala Asp Thr Thr His Ser Thr Val Leu

Gly Leu Tyr Asn Thr Leu Asn Pro Glu Ala Ser Ala Ser Pro Cys Cys 55

Val Pro Gln Asp Leu Glu Pro Leu Thr Ile Leu Tyr Tyr Val Gly Arg 70 75

Thr Pro Lys Val Glu Gln Leu Ser Asn Met Val Val Lys Ser Cys Lys 85 90

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<210> 43

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<213> Gallus gallus

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Lys Trp Ile His Glu Pro Lys Gly Tyr Met Ala Asn Phe Cys Met Gly
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Pro Cys Pro Tyr Ile Trp Ser Ala Asp Thr Gln Tyr Thr Lys Val Leu 35 40 45

Ala Leu Tyr Asn Gln His Asn Pro Gly Ala Ser Ala Ala Pro Cys Cys 50 55 60

Val Pro Gln Thr Leu Asp Pro Leu Pro Ile Ile Tyr Tyr Val Gly Arg
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Asn Val Arg Val Glu Gln Leu Ser Asn Met Val Val Arg Ala Cys Lys 85 90 95

Cys Ser

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<213> Xenopus laevis

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Asn Cys Pro Tyr Ile Trp Ser Met Asp Thr Gln Tyr Ser Lys Val Leu 35 40 45

Ser Leu Tyr Asn Gln Asn Asn Pro Gly Ala Ser Ile Ser Pro Cys Cys 50 55 60

Val Pro Asp Val Leu Glu Pro Leu Pro Ile Ile Tyr Tyr Val Gly Arg 65 70 75 80

Thr Ala Lys Val Glu Gln Leu Ser Asn Met Val Val Arg Ser Cys Asn 85 90 95

Cys Ser

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Lys Cys Pro Phe Pro Leu Ala Asp His Phe Asn Ser Thr Asn His Ala 35 40 45

Val Val Gln Thr Leu Val Asn Asn Met Asn Pro Gly Lys Val Pro Lys 50 55 60

Ala Cys Cys Val Pro Thr Gln Leu Asp Ser Val Ala Met Leu Tyr Leu 65 70 75 80

Asn Asp Gln Ser Thr Val Val Leu Lys Asn Tyr Gln Glu Met Thr Val 85 90 95

Val Gly Cys Gly Cys Arg

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<213> Xenopus laevis

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20 25 30

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Ile Leu Gln Thr Leu Val His Ser Ile Glu Pro Glu Asp Ile Pro Leu

Pro Cys Cys Val Pro Thr Lys Met Ser Pro Ile Ser Met Leu Phe Tyr

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Glu Cys Ser Phe Pro Leu Asn Ala His Met Asn Ala Thr Asn His Ala 40

Ile Val Gln Thr Leu Val His Leu Met Asn Pro Glu Tyr Val Pro Lys 55

Pro Cys Cys Ala Pro Thr Lys Leu Asn Ala Ile Ser Val Leu Tyr Phe 75

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Arg Ala Cys Gly Cys His 100

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Ile Val Gln Thr Leu Val Asn Ser Val Asn Ser Lys Ile Pro Lys Ala 55

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Glu Asn Glu Lys Val Val Leu Lys Asn Tyr Gln Asp Met Val Val Glu 90

Gly Cys Gly Cys Arg 100

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Glu Trp Ile Ile Ser Pro Lys Ser Phe Asp Ala Tyr Tyr Cys Ser Gly

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Thr Ile Gln Ser Ile Val Arg Ala Val Gly Val Val Pro Gly Ile Pro 55

Glu Pro Cys Cys Val Pro Glu Lys Met Ser Ser Leu Ser Ile Leu Phe 75

Phe Asp Glu Asn Lys Asn Val Val Leu Lys Val Tyr Pro Asn Met Thr 85 90

Val Glu Ser Cys Ala Cys Arg 100

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60

Ile Val Gln Thr Leu Val His Leu Met Phe Pro Asp His Val Pro Lys

Pro Cys Cys Ala Pro Thr Lys Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val 90 Arg Ser Cys Gly Cys His 100 <210> 53 <211> 102 <212> PRT <213> Homo sapiens <220> <223> BMP-6 <400> 53 Cys Arg Lys His Glu Leu Tyr Val Ser Phe Gln Asp Leu Gly Trp Gln Asp Trp Ile Ile Ala Pro Lys Gly Tyr Ala Ala Asn Tyr Cys Asp Gly Glu Cys Ser Phe Pro Leu Asn Ala His Met Asn Ala Thr Asn His Ala Ile Val Gln Thr Leu Val His Leu Met Asn Pro Glu Tyr Val Pro Lys Pro Cys Cys Ala Pro Thr Lys Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp Asp Asn Ser Asn Val Glu Leu Lys Lys Tyr Arg Asn Met Val Val Arg Ala Cys Gly Cys His 100 <210> 54 <211> 103 <212> PRT <213> Gallus gallus

<220>

<223> DORSALIN

<400> 54

Cys Arg Arg Thr Ser Leu His Val Asn Phe Lys Glu Ile Gly Trp Asp Ser Trp Ile Ile Ala Pro Lys Asp Tyr Glu Ala Phe Glu Cys Lys Gly 25 Gly Cys Phe Phe Pro Leu Thr Asp Asn Val Thr Pro Thr Lys His Ala Ile Val Gln Thr Leu Val His Leu Gln Asn Pro Lys Lys Ala Ser Lys 55 Ala Cys Cys Val Pro Thr Lys Leu Asp Ala Ile Ser Ile Leu Tyr Lys Asp Asp Ala Gly Val Pro Thr Leu Ile Tyr Asn Tyr Glu Gly Met Lys Val Ala Glu Cys Gly Cys Arg 100 <210> 55 <211> 102 <212> PRT <213> Homo sapiens <220> <223> OP-1 <400> 55 Cys Lys Lys His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Gln Asp Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly Glu Cys Ala Phe Pro Leu Asn Ser Tyr Met Asn Ala Thr Asn His Ala 40 Ile Val Gln Thr Leu Val His Phe Ile Asn Pro Glu Thr Val Pro Lys 55 60 Pro Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile Ser Val Leu Tyr Phe 70 75 Asp Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Arg Asn Met Val Val

Page 28

85 90 95

Arg Ala Cys Gly Cys His

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<211> 102

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<213> Homo sapiens

<220>

<223> OP-2

<400> 56

Cys Arg Arg His Glu Leu Tyr Val Ser Phe Gln Asp Leu Gly Trp Leu 1 5 10 15

Asp Trp Val Ile Ala Pro Gln Gly Tyr Ser Ala Tyr Tyr Cys Glu Gly 20 25 30

Glu Cys Ser Phe Pro Leu Asp Ser Cys Met Asn Ala Thr Asn His Ala 35 40 45

Ile Leu Gln Ser Leu Val His Leu Met Lys Pro Asn Ala Val Pro Lys 50 55 60

Ala Cys Cys Ala Pro Thr Lys Leu Ser Ala Thr Ser Val Leu Tyr Tyr 65 70 75 80

Asp Ser Ser Asn Asn Val Ile Leu Arg Lys His Arg Asn Met Val Val
85 90 95

Lys Ala Cys Gly Cys His 100

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Asp Ser Val Ile Ala Pro Gln Gly Tyr Ser Ala Tyr Tyr Cys Ala Gly

Glu Cys Ile Tyr Pro Leu Asn Ser Cys Met Asn Ser Thr Asn His Ala

Thr Met Gln Ala Leu Val His Leu Met Lys Pro Asp Ile Ile Pro Lys 55

Val Cys Cys Val Pro Thr Glu Leu Ser Ala Ile Ser Leu Leu Tyr Tyr

Asp Arg Asn Asn Val Ile Leu Arg Arg Glu Arg Asn Met Val Val 90

Gln Ala Cys Gly Cys His 100

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Cys Arg Thr Arg Arg Leu His Val Ser Phe Arg Glu Val Gly Trp His

Arg Trp Val Ile Ala Pro Arg Gly Phe Leu Ala Asn Phe Cys Gln Gly

Thr Cys Ala Leu Pro Glu Thr Leu Arg Gly Pro Gly Gly Pro Pro Ala

Leu Asn His Ala Val Leu Arg Ala Leu Met His Ala Ala Pro Thr 55

Pro Gly Ala Gly Ser Pro Cys Cys Val Pro Glu Arg Leu Ser Pro Ile 75

Ser Val Leu Phe Phe Asp Asn Ser Asp Asn Val Val Leu Arg His Tyr 85 90

Glu Asp Met Val Val Asp Glu Cys Gly Cys Arg 100 105

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Cys His Arg His Gln Leu Phe Ile Asn Phe Gln Asp Leu Gly Trp His
Lys Trp Val Ile Ala Pro Lys Gly Phe Met Ala Asn Tyr Cys His Gly
                                25
Glu Cys Pro Phe Ser Met Thr Thr Tyr Leu Asn Ser Ser Asn Tyr Ala
                            40
Phe Met Gln Ala Leu Met His Met Ala Asp Pro Lys Val Pro Lys Ala
Val Cys Val Pro Thr Lys Leu Ser Pro Ile Ser Met Leu Tyr Gln Asp
Ser Asp Lys Asn Val Ile Leu Arg His Tyr Glu Asp Met Val Val Asp
Glu Cys Gly Cys Gly
          100
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Cys Glu Leu His Asp Phe Arg Leu Ser Phe Ser Gln Leu Lys Trp Asp
Asn Trp Ile Val Ala Pro His Arg Tyr Asn Pro Arg Tyr Cys Lys Gly
Asp Cys Pro Arg Ala Val Arg His Arg Tyr Gly Ser Pro Val His Thr
                            40
                                               4.5
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Met Val Gln Asn Ile Ile Tyr Glu Lys Leu Asp Pro Ser Val Pro Arg Pro Ser Cys Val Pro Gly Lys Tyr Ser Pro Leu Ser Val Leu Thr Ile Glu Pro Asp Gly Ser Ile Ala Tyr Lys Glu Tyr Glu Asp Met Ile Ala 90 Thr Arg Cys Thr Cys Arg 100 <210> 61 <211> 105 <212> PRT <213> Homo sapiens <220> <223> INHIBIN-Alpha <400> 61 Cys His Arg Val Ala Leu Asn Ile Ser Phe Gln Glu Leu Gly Trp Glu Arg Trp Ile Val Tyr Pro Pro Ser Phe Ile Phe His Tyr Cys His Gly Gly Cys Gly Leu His Ile Pro Pro Asn Leu Ser Leu Pro Val Pro Gly Ala Pro Pro Thr Pro Ala Gln Pro Tyr Ser Leu Leu Pro Gly Ala Gln 55 Pro Cys Cys Ala Ala Leu Pro Gly Thr Met Arg Pro Leu His Val Arg Thr Thr Ser Asp Gly Gly Tyr Ser Phe Lys Tyr Glu Thr Val Pro Asn 90 Leu Leu Thr Gln His Cys Ala Cys Ile 100 <210> 62 <211> 106

<212> PRT

<213> Bos taurus

<220>

<223> INHIBIN-BetaA

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Cys Cys Lys Lys Gln Phe Phe Val Ser Phe Lys Asp Ile Gly Trp Asn 1 5 10 15

Asp Trp Ile Ile Ala Pro Ser Gly Tyr His Ala Asn Tyr Cys Glu Gly 20 25 30

Glu Cys Pro Ser His Ile Ala Gly Thr Ser Gly Ser Ser Leu Ser Phe 35 40 45

His Ser Thr Val Ile Asn His Tyr Arg Met Arg Gly His Ser Pro Phe 50 55 60

Ala Asn Leu Lys Ser Cys Cys Val Pro Thr Lys Leu Arg Pro Met Ser 65 70 75 80

Met Leu Tyr Tyr Asp Asp Gly Gln Asn Ile Ile Lys Lys Asp Ile Gln 85 90 95

Asn Met Ile Val Glu Glu Cys Gly Cys Ser 100 105

<210> 63

<211> 106

<212> PRT

<213> Homo sapiens

<220>

<223> INHIBIN-Betab

<400> 63

Cys Cys Lys Lys Gln Phe Phe Val Ser Phe Lys Asp Ile Gly Trp Asn 1 $$ 5 $$ 10 $$ 15

Asp Trp Ile Ile Ala Pro Ser Gly Tyr His Ala Asn Tyr Cys Glu Gly
20 25 30

Glu Cys Pro Ser His Ile Ala Gly Thr Ser Gly Ser Ser Leu Ser Phe 35 40 45

His Ser Thr Val Ile Asn His Tyr Arg Met Arg Gly His Ser Pro Phe 50 60

Ala Asn Leu Lys Ser Cys Cys Val Pro Thr Lys Leu Arg Pro Met Ser

Met Leu Tyr Tyr Asp Asp Gly Gln Asn Ile Ile Lys Lys Asp Ile Gln 85 90 95

Asn Met Ile Val Glu Glu Cys Gly Cys Ser 100 105

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<213> Artificial Sequence

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<222> (12)..(12)

<223> Xaa12 can be Arg or Lys

<220>

<221> misc_feature

<222> (26)..(26)

<223> Xaa26 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val

<220>

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<222> (31)..(31)

<223> Xaa31 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val

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<222> (33)..(33)

<223> Xaa33 can be Ala, Gly, Pro, Ser or Thr

<220>

<221> misc_feature

<222> (37)..(37)

<223> Xaa37 can be Ile, Leu lys, Met or Val

<220>

<221> misc_feature

<222> (40)..(40)

<223> Xaa40 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val

<220>

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<222> (44)..(44)

<223> Xaa44 can be His, Phe, Trp or Tyr

<220>

<221> misc feature

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<223> Xaa46 can be Arg or Lys
<220>
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<223> Xaa49 can be Ala, Gly, Pro, Ser or Thr
<220>
<221> misc feature
<222> (53)..(54)
<223> Xaa53 can be Arg, Asn, Asp, Gln, Glu, His, Lys, Ser or Thr;
       Xaa54 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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<223>
      Xaa57 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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      (61)..(61)
<223> Xaa61 can be Ala, Gly, Pro, Ser or Thr
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      (68)..(68)
<222>
      Xaa68 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
      Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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<223> Xaa73 can be Ala, Gly, Pro, Ser or Thr
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      (75)..(75)
<223> Xaa75 can be Ile, Leu, Met or Val
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      (81)..(82)
<223> Xaa81 can be Arg, Asn, Asp, Gln, Glu, His, Lys, Ser or Thr;
      Xaa82 can be Ala, Gly, Pro, Ser, or Thr
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<222> (91)..(91)
<223> Xaa91 can be any Ile or Val
<220>
<221> misc_feature
<222>
      (93)..(93)
<223> Xaa93 can be Arg or Lys
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Cys Cys Val Arg Pro Leu Tyr Ile Asp Phe Arg Xaa Asp Leu Gly Trp Lys Trp Ile His Glu Pro Lys Gly Tyr Xaa Ala Asn Phe Cys Xaa Gly 25 Xaa Cys Pro Tyr Xaa Trp Ser Xaa Asp Thr Gln Xaa Ser Xaa Val Leu 40 Xaa Leu Tyr Asn Xaa Xaa Asn Pro Xaa Ala Ser Ala Xaa Pro Cys Cys 55 Val Pro Gln Xaa Leu Glu Pro Leu Xaa Ile Xaa Tyr Tyr Val Gly Arg 75 70 Xaa Xaa Lys Val Glu Gln Leu Ser Asn Met Xaa Val Xaa Ser Cys Lys 90 Cys Ser <210> 65 <211> 104 <212> PRT <213> Artificial Sequence <220> <223> Description of Artificial Sequence: VG/DPP SUBGROUP SEQUENCE PATTERN <220> <221> misc_feature <222> (2)..(5) <223> Xaa2 can be Arg or Lys; Xaa3 can be Arg or Lys; Xaa4 and Xaa5 independently can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc_feature <222> (9)..(9) <223> Xaa9 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc_feature <222> (11)..(11) <223> Xaa11 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc feature <222> (13)..(13) <223> Xaa13 can be Ile, Leu, Met, or Val <220>

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      (16)..(16)
<223> Xaa16 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
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<222> (23)..(23)
<223> Xaa23 can be Arg, Gln, Glu or Lys
<220>
<221> misc_feature
<222>
      (26)..(26)
<223>
      Xaa26 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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<222>
      (28)..(28)
<223> Xaa28 can be Phe, Trp or Tyr
<220>
<221> misc_feature
<222>
      (31)..(31)
<223> Xaa31 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
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      (33)..(33)
<223> Xaa33 can be Asp or Glu
<220>
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      (35)..(35)
<223> Xaa35 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
      Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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      (39)..(42)
<223> Xaa39, Xaa40 and Xaa41 independently can be Ala, Arg, Asn,
      Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro,
      Ser, Thr, Trp, Tyr or Val; Xaa42 can be Leu or Met
<220>
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<222> (44)..(44)
<223> Xaa44 can be Ala, Gly, Pro, Ser or Thr
<220>
<221> misc feature
<222> (50)..(50)
<223> Xaa50 can be Ile or Val
<220>
<221> misc feature
      (55)..(57)
<222>
<223> Xaa55 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr;
      Xaa56 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile,
      Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa57 can
      be Ile, Leu, Met or Val
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      (58)..(60)
<223> Xaa58 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr;
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       Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr,
       Trp, Tyr, Val or a peptide bond
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      (61)..(63)
      Xaa61 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr;
<223>
       Xaa62 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile,
       Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa63
       can be Ile or Val
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      (66)..(66)
<223> Xaa66 can be Ala Gly, Pro, Ser or Thr
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      (69)..(69)
      Xaa69 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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<222>
      (72)..(72)
<223> Xaa72 can be Arg, Gln, Glu or Lys
<220>
<221> misc_feature
<222> (74)..(74)
<223> Xaa74 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
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<222> (76)..(76)
<223> Xaa76 can be Ile or Val
<220>
<221> misc feature
<222> (78)..(78)
<223> Xaa78 can be Ile, Leu, Met or Val
<220>
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<222> (81)..(81)
<223> Xaa81 can be Cys, Ile, Leu, Met, Phe, Trp, Tyr or Val
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<222> (83)..(85)
      Xaa83 can be Asn, Asp or Glu; Xaa84 can be Arg, Asn, Asp,
      Glu, Gln, His, Lys, Ser or Thr; Xaa85 can be Ala, Arg,
      Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe,
      Pro, Ser, Thr, Trp, Tyr, Val or a peptide bond
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<223> Xaa86 and Xaa87 independently can be Arg, Asn, Asp, Glu,
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<222> (89)..(89)
<223> Xaa89 can Ile or Val
<220>
<221> misc feature
<222>
      (91)..(92)
<223> Xaa91 can be Arg or Lys; Xaa92 can be Arg, Asn, Asp,
      Glu, Gln, His, Lys, Ser or Thr
<220>
<221> misc_feature
<222>
      (94)..(95)
<223> Xaa94 can be Arg, Gln, Glu or Lys; Xaa95 can be Asn or Asp
<220>
<221> misc_feature
<222>
      (97)..(97)
      Xaa97 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
<223>
      Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
<220>
<221> misc_feature
<222> (99)..(100)
<223> Xaa99 can be Arg, Gln, Glu or Lys; Xaa100 can be Ala Gly,
      Pro, Ser or Thr
<220>
<221> misc_feature
      (104)..(104)
<223> Xaa104 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
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Cys Xaa Xaa Xaa Leu Tyr Val Xaa Phe Xaa Asp Xaa Gly Trp Xaa
               5
                                                        15
                                    10
Asp Trp Ile Ile Ala Pro Xaa Gly Tyr Xaa Ala Xaa Tyr Cys Xaa Gly
           20
                                25
                                                    30
Xaa Cys Xaa Phe Pro Leu Xaa Xaa Xaa Asn Xaa Thr Asn His Ala
       35
                            40
Ile Xaa Gln Thr Leu Val Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Aaa Pro
   50
                       55
                                            60
Lys Xaa Cys Cys Xaa Pro Thr Xaa Leu Xaa Ala Xaa Ser Xaa Leu Tyr
65
                    70
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Xaa Asp Xaa Xaa Xaa Xaa Val Xaa Leu Xaa Xaa Tyr Xaa Xaa Met 85 90 Xaa Val Xaa Xaa Cys Gly Cys Xaa 100 <210> 66 <211> 107 <212> PRT <213> Artificial Sequence <220> <223> Description of Artificial Sequence: GDF SUBGROUP PATTERN <220> <221> misc_feature <222> (2)..(3) Xaa2 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr; <223> Xaa3 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val <220> <221> misc feature <222> (4)..(5) <223> Xaa4 and Xaa5 independently can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc_feature <222> (6)..(8) <223> Xaa6 can be Cys, Ile, Leu, Met, Phe, Trp, Tyr or Val; Xaa7 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa8 can be Ile, Leu, Met or Val <220> <221> misc feature (9)..(9) <222> <223> Xaa9 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Th <220> <221> misc_feature <222> (11)..(14) <223> Xaall and Xaal2 independently can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr; Xaal3 can be Ile, Leu, Met or Val; Xaa14 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val <220> <221> misc feature

<220>

<222>

<221> misc feature

(16)..(17)

His, Lys, Ser or Thr

<223> Xaa16 and Xaa17 independently can be Arg, Asn, Asp, Glu, Gln,

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<223> Xaa19 and Xaa20 independently can be Ile or Val
<220>
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<222>
       (23)..(25)
       Xaa23 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr;
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       Xaa24 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile,
       Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa25
       can be Phe, Trp or Tyr
<220>
<221> misc_feature
<222>
       (26)..(29)
       Xaa26 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile,
<223>
       Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa27 can
       be Ala Gly, Pro, Ser or Thr; Xaa28 can be Arg, Asn, Asp, Glu,
       Gln, His, Lys, Ser or Thr; Xaa29 can be Phe, Trp or Tyr
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      (31)..(31)
<223> Xaa31 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
<220>
<221> misc_feature
<222> (33)..(33)
<223> Xaa33 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
<220>
<221> misc feature
<222>
      (35)..(37)
<223> Xaa35 can be Ala, Gly, Pro, Ser or Thr; Xaa36 can be Ala, Arg,
       Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe,
       Pro, Ser, Thr, Trp, Tyr or Val; Xaa37 can be Ala, Gly, Pro,
       Ser or Thr
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<222> (38)..(39)
<223> Xaa38 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile,
       Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val;
       can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
<220>
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      (40)..(42)
<222>
<223> Xaa40 to Xaa42 independently can be Ala, Arg, Asn, Asp,
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       Thr, Trp, Tyr or Val
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      (43)..(46)
<223> Xaa43 to Xaa46 independently can be Ala, Arg, Asn,
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       Ser, Thr, Trp, Tyr, Val or a peptide bond
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- <222> (47)..(48)
- <223> Xaa47 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa48 can be Ala, Gly, Pro, Ser or Thr
- <220>
- <221> misc_feature
- <222> (49)..(49)
- <223> Xaa49 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
- <220>
- <221> misc feature
- <222> (50)..(53)
- <223> Xaa50 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa51 can be His, Phe, Trp or Tyr; Xaa52 can be Ala, Gly, Pro, Ser or Thr; Xaa53 can be Cys, Ile, Leu, Met, Phe, Trp, Tyr or Val
- <220>
- <221> misc_feature
- <222> (54)..(55)
- <223> Xaa54 can be Ile, Leu, Met or Val; Xaa55 can be Arg, Gln, Glu or Lys; Xaa56 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa57 and Xaa58 independently can be Ile, Leu, Met or Val
- <220>
- <221> misc_feature
- <222> (59)..(62)
- <223> Xaa59 can be His, Phe, Trp or Tyr; Xaa60, Xaa61 and Xaa62 independently can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
- <220>
- <221> misc_feature
- <222> (63)..(64)
- <223> Xaa63 and Xaa64 independently can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr, Val or a peptide bond
- <220>
- <221> misc_feature
- <222> (66)..(69)
- <223> Xaa66 and Xaa67 independently can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa68 can be Ala, Gly, Pro, Ser or Thr; Xaa69 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
- <220>
- <221> misc feature
- <222> (70)..(71)
- <223> Xaa70 can be Ala, Gly, Pro, Ser or Thr; Xaa71 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
- <220>
- <221> misc feature
- <222> (75)..(77)
- <223> Xaa75 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,

Phe, Trp, Tyr or Val <220> <221> misc_feature <222> (80)..(80) <223> Xaa80 can be Ile, Leu, Met or Val <220> <221> misc feature <222> (82)..(82) <223> Xaa82 can be Ile, Leu, Met or Val <220> <221> misc feature <222> (84)..(86) <223> Xaa84 and Xaa85 independently can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa86 can be Asp or Glu <220> <221> misc feature <222> (87)..(88) Xaa87 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, <223> Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa88 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc_feature <222> (89)..(91) Xaa89 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, <223> Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa90 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr; Xaa 91 is Ile or Val <220> <221> misc feature <222> (92)..(94) <223> Xaa92 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa93 can be Cys, Ile, Leu, Met, Phe, Trp, Tyr or Val; Xaa94 can be Arg or Lys <220> <221> misc feature <222> (95)..(95) <223> Xaa95 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc feature <222> (100)..(103) Xaa100 can be Ile or Val; Xaa101 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa102 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr; Xaa103 can be Arg, Gln, Glu or Lys <220> <221> misc feature <222> (105)..(105)

Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa76 can be Arg, or Lys; Xaa77 can be Cys, Ile, Leu, Met,

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<223> Xaa105 can be Ala, Gly, Pro, Ser or Thr
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<222> (107)..(107)
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      Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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Cys Xaa Xaa Xaa Xaa Xaa Xaa Xaa Phe Xaa Xaa Xaa Xaa Trp Xaa
                              10
Xaa Trp Xaa Xaa Ala Pro Xaa Xaa Xaa Xaa Xaa Xaa Cys Xaa Gly
          20
                          25
40
Pro Xaa Xaa Xaa Xaa Xaa Cys Val Pro Xaa Xaa Ser Pro Xaa
85
                              90
Glu Asp Met Xaa Xaa Xaa Cys Xaa Cys Xaa
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     Xaa3 is Arg or Lys
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     (4)..(6)
     Xaa4 and Xaa5 independently can be Ala, Arg, Asn, Asp,
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     Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro,
     Ser, Thr, Trp, Tyr or Val; Xaa6 can be Cys, Ile, Leu,
     Met, Phe, Trp, Tyr or Val
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      (7)..(9)
<223> Xaa7 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val;
       Xaa8 can be Ile or Val; Xaa9 can be Arg, Asn, Asp, Glu,
       Gln, His, Lys, Ser or Thr
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<222>
      (11)..(13)
<223>
      Xaa11 can be Arg, Gln, Glu or Lys; Xaa12 can be Ala, Arg,
       Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met,
       Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa13 can be Ile,
       Leu, Met or Val
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      Xaal6 can be Asn, Asp or Glu; Xaal7 can be Arg, Asn, Asp,
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       Glu, Gln, His, Lys, Ser or Thr
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      Xaa20 can be Ile or Val; Xaa21 can be Ala, Arg, Asn, Asp,
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       Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser,
       Thr, Trp, Tyr or Val
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<223> Xaa23 and Xaa24 independently can be Ala, Gly, Pro, Ser or
       Thr; Xaa25 can be Phe, Tr or Tyr; Xaa26 and Xaa27
       independently can beAla, Arg, Asn, Asp, Cys, Glu, Gln, Gly,
      His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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<223> Xaa28 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
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<221> misc feature
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<223> Xaa31 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr
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<223> Xaa33 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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      (35)..(38)
<223> Xaa35 can be Ala, Gly, Pro, Ser or Thr; Xaa36 can be Ala,
      Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met,
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Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa37 can be His, Phe,

Trp or Tyr; Xaa38 can be Ile, Leu, Met or Val

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<222>
<223>
       Xaa39 and Xaa40 independently can be Ala, Gly, Pro, Ser or
       Thr; Xaa41 and Xaa42 independently can be Ala, Arg, Asn, Asp,
       Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser,
       Thr, Trp, Tyr or Val; Xaa43 can be Ala, Gly, Pro, Ser or Thr
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       Xaa44 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
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       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val;
       Xaa45 can be Ala, Gly, Pro, Ser or Thr
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       Xaa46 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val;
       Xaa47 can be Ala, Gly, Pro, Ser or Thr
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      (48)..(51)
<223>
       Xaa48 and Xaa49 independently can be Ala, Arg, Asn, Asp,
       Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser,
       Thr, Trp, Tyr or Val; Xaa50 and Xaa51 independently can be
       Ala, Gly, Pro, Ser or Thr
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      (52)..(55)
<223> Xaa52 to Xaa54 independently can be Ala, Arg, Asn,
       Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro,
       Ser, Thr, Trp, Tyr or Val; Xaa55 can be Arg, Asn, Asp, Glu,
       Gln, His, Lys, Ser or Thr
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      (56)..(59)
      Xaa56 to Xaa59 indepedently can be Ala, Arg, Asn,
<223>
       Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro,
       Ser, Thr, Trp, Tyr or Val
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<221> misc_feature
<222> (60)..(63)
<223>
       Xaa60 to Xaa63 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln,
       Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr,
       Val or a peptide bond
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<221> misc feature
<222>
      (64)..(65)
<223> Xaa64 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val;
       Xaa65 can be Ala, Gly, Pro, Ser or Thr
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      (66)..(69)
<223> Xaa66 to Xaa67 independently can be Ala, Arg, Asn, Asp,
       Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser,
       Thr, Trp, Tyr or Val; Xaa68 can be Arg, Asn, Asp, Glu, Gln,
       His, Lys, Ser or Thr; Xaa69 can be Ala, Gly, Pro, Ser or Thr
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      (72)..(72)
<222>
<223> Xaa72 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile,
       Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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<222>
      (73)..(74)
      Xaa73 and Xaa74 independently can be Ala, Arg, Asn, Asp,
<223>
       Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser,
       Thr, Trp, Tyr or Val
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      (76)..(80)
      Xaa76 can be Ala, Gly, Pro, Ser or Thr; Xaa77 can be Arg,
<223>
      Asn, Asp, Glu, Gln, His, Lys, Ser or Thr; Xaa78 can be Leu
       or Met; Xaa79 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser
       or Thr; Xaa80 can be Ala, Gly, Pro, Ser or Thr
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<221> misc_feature
<222>
      (81)..(83)
      Xaa81 can be Leu or Met; Xaa82 can be Arg, Asn, Asp, Glu,
      Gln, His, Lys, Ser or Thr; Xaa83 can be Ile, Leu, Met or Val
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<222>
<223>
      Xaa84 to Xaa86 independently can be Ala, Arg, Asn, Asp,
      Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser,
       Thr, Trp, Tyr or Val; Xaa87 can be Arg, Asn, Asp, Glu, Gln,
      His, Lys, Ser or Thr
<220>
<221> misc feature
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      (89)..(89)
      Xaa89 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val
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      Xaa90 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His,
       Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr, Val or
       a peptide bond
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<222> (91)..(93)

- <223> Xaa91 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa92 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr; Xaa93 can be Cys, Ile, Leu, Met, Phe, Trp, Tyr or Val <220> <221> misc feature <222> (94)..(97) Xaa94 to Xaa95 independently can be Ala, Arg, Asn, Asp, Cys, <223> Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa96 can be Arg, Gln, Glu or Lys; Xaa97 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc feature <222> (98)..(99) Xaa98 can be Ile or Val; Xaa99 can be Ala, Arg, Asn, Asp, <223> Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val <220> <221> misc feature (101) .. (104) <222> Xaa101 can be Leu or Met; Xaa102 can be Ile, Leu, Met or <223> Val; Xaa103 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val; Xaa104 can be Gln or Glu <220> <221> misc_feature <222> (105)..(105) <223> Xaa105 can be Arg, Asn, Asp, Glu, Gln, His, Lys, Ser or Thr <220> <221> misc_feature <222> (107)..(107) <223> Xaa107 can be Ala or Gly <220> <221> misc feature <222> (109)..(109) Xaa109 can be Ala, Arg, Asn, Asp, Cys, Glu, Gln, Gly, His, Ile, Leu, Lys, Met, Phe, Pro, Ser, Thr, Trp, Tyr or Val <400> Cys Xaa Xaa Xaa Xaa Xaa Xaa Xaa Phe Xaa Xaa Gly Trp Xaa 5 15 Xaa Trp Ile Xaa Xaa Pro Xaa Xaa Xaa Xaa Xaa Tyr Cys Xaa Gly

Xaa 65	Xaa	Xaa	Xaa	Xaa	Cys 70	Cys	Xaa	Xaa	Xaa	Pro 75	Xaa	Xaa	Хаа	Xaa	Xaa 80
Xaa	Xaa	Xaa	Xaa	Xaa 85	Xaa	Xaa	Asp	Xaa	Xaa 90	Xaa	Xaa	Xaa	Xaa	Xaa 95	Xaa
Xaa	Xaa	Xaa	Asn 100	Xaa	Xaa	Xaa	Xaa	Xaa 105	Сув	Xaa	Сув	Xaa			
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<400	O > (68													
Ser 1	Thr	Gly	Ser	Lys 5	Gln	Arg	Ser	Gln	Asn 10	Arg	Ser	Lys	Thr	Pro 15	Lys
Asn	Gln	Glu	Ala 20	Leu	Arg	Met	Ala	Asn 25	Val	Ala	Glu	Asn	Ser 30	Ser	Ser
Asp	Gln	Arg 35	Gln	Ala	Cys	Lys	Lys 40	His	Glu	Leu	Tyr	Val 45	Ser	Phe	Arg
Asp	Leu 50	Gly	Trp	Gln	Asp	Trp 55	Ile	Ile	Ala	Pro	Glu 60	Gly	Tyr	Ala	Ala
Tyr 65	Tyr	Cys	Glu	Gly	Glu 70	Cys	Ala	Phe	Pro	Leu 75	Asn	Ser	Tyr	Met	Asn 80
Ala	Thr	Asn	His	Ala 85	Ile	Val	Gln	Thr	Leu 90	Val	His	Phe	Ile	Asn 95	Pro
Glu	Thr	Val	Pro 100	Lys	Pro	Сув	Cys	Ala 105	Pro	Thr	Gln	Leu	Asn 110	Ala	Ile
Ser	Val	Leu 115	Tyr	Phe	Asp	Asp	Ser 120	Ser	Asn	Val	Ile	Leu 125	Lys	Lys	Tyr
Glu	Asp 130	Met	Val	Val	Glu	Ala 135	Сув	Gly	Cys	Arg					

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<213> Homo sapiens
<223> Trypsin truncated H2223 mutant
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Met Ala Asn Val Ala Glu Asn Ser Ser Ser Asp Gln Arg Gln Ala Cys
                                    10
Lys Lys His Glu Leu Tyr Val Ser Phe Arg Asp Leu Gly Trp Gln Asp
            20
                                25
Trp Ile Ile Ala Pro Glu Gly Tyr Ala Ala Tyr Tyr Cys Glu Gly Glu
                            40
Cys Ala Phe Pro Leu Asn Ser Tyr Met Asn Ala Thr Asn His Ala Ile
    50
                       55
Val Gln Thr Leu Val His Phe Ile Asn Pro Glu Thr Val Pro Lys Pro
                    70
Cys Cys Ala Pro Thr Gln Leu Asn Ala Ile Ser Val Leu Tyr Phe Asp
Asp Ser Ser Asn Val Ile Leu Lys Lys Tyr Glu Asp Met Val Val Glu
Ala Cys Gly Cys Arg
       115
<210> 70
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Ala Pro Thr Gln Leu Ser Ala Ile Ser Val Leu
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<223> Amino acid sequence encoded by Primer #1
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Ala Pro Thr Gln Leu Ser Ala Ile Ser Val Leu
<210> 72
<211> 43
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ctatctgcag ccacaagctt cgaccaccat gtcttcgtat ttc
                                                                      43
<210> 73
<211>
      43
<212> DNA
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g aaa tac gaa gac atg gtg gtc gaa gct tgt ggc tgc aga tag
                                                                     43
 Lys Tyr Glu Asp Met Val Val Glu Ala Cys Gly Cys Arg
<210> 74
<211> 13
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Lys Tyr Glu Asp Met Val Val Glu Ala Cys Gly Cys Arg
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<210> 75

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<211> 44
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      between the T7 promoter, at the XbaI site, and the
       ATG codon
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                                                                      44
tctagaataa ttttgtttaa cctttaagaa ggagatatac gatg
<210> 76
<211> 19
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Primer #3
<400> 76
taatacgact cactatagg
                                                                      19
<210> 77
<211> 18
<212> DNA
<213> Artificial Sequence
<220>
<223> Description of Artificial Sequence: Primer #4
<400> 77
                                                                     18
gctgagctgc gtgggcgc
<210> 78
<211> 18
<212> DNA
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<223> Description of Artificial Sequence: complement of Primer #4
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<221> CDS
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gcg ccc acg cag ctc agc
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Ala Pro Thr Gln Leu Ser
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<210> 79
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Ala Pro Thr Gln Leu Ser
<210> 80
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ggatcctatc tgcagccaca agc
<210> 81
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<223> Description of Artificial Sequence: complement of Primer #5
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<400> 81
gct tgt ggc tgc aga tag gatcc
                                                                     23
Ala Cys Gly Cys Arg
<210> 82
<211> 5
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Ala Cys Gly Cys Arg
<210> 83
<211> 102
<212> PRT
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<213> Homo sapiens

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<223> CDMP-1/GDF-5
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Cys Ser Arg Lys Ala Leu His Val Asn Phe Lys Asp Met Gly Trp Asp
Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Phe His Cys Glu Gly
Leu Cys Glu Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala
Val Ile Gln Thr Leu Met Asn Ser Met Asp Pro Glu Ser Thr Pro Pro
Thr Cys Cys Val Pro Thr Arg Leu Ser Pro Ile Ser Ile Leu Phe Ile
                                         75
Asp Ser Ala Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val
Glu Ser Cys Gly Cys Arg
            100
<210> 84
<211> 102
<212> PRT
<213> Homo sapiens
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<223> CDMP-2/GDF-6
<400> 84
Cys Ser Lys Lys Pro Leu His Val Asn Phe Lys Glu Leu Gly Trp Asp
Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Tyr His Cys Glu Gly
            20
Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala
                            40
Ile Ile Gln Thr Leu Met Asn Ser Met Asp Pro Gly Ser Thr Pro Pro
   50
                        55
                                             60
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Ser Cys Cys Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile

Asp Ala Gly Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val 85 90 95

Glu Ser Cys Gly Cys Arg 100

70

<210> 85

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<212> PRT

<213> Mus musculus

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<223> GDF-6

<400> 85

Cys Ser Arg Lys Pro Leu His Val Asn Phe Lys Glu Leu Gly Trp Asp 1 5 10 15

Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Tyr His Cys Glu Gly
20 25 30

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala 35 40- 45

Ile Ile Gln Thr Leu Met Asn Ser Met Asp Pro Gly Ser Thr Pro Pro 50 55 60

Ser Cys Cys Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile 65 70 75 80

Asp Ala Gly Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val 85 90 95

Glu Ser Cys Gly Cys Arg 100

<210> 86

<211> 102

<212> PRT

<213> Bos taurus

<220>

<223> CDMP-2

<400> 86

Cys Ser Lys Lys Pro Leu His Val Asn Phe Lys Glu Leu Gly Trp Asp

Asp Trp Ile Ile Ala Pro Leu Glu Tyr Glu Ala Tyr His Cys Glu Gly

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala

Ile Ile Gln Thr Leu Met Asn Ser Met Asp Pro Gly Ser Thr Pro Pro

Ser Cys Cys Val Pro Thr Lys Leu Thr Pro Ile Ser Ile Leu Tyr Ile

Asp Ala Gly Asn Asn Val Val Tyr Asn Glu Tyr Glu Glu Met Val Val

Glu Ser Cys Gly Cys Arg 100

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<223> GDF-7

<400> 87

Cys Ser Arg Lys Ser Leu His Val Asp Phe Lys Glu Leu Gly Trp Asp

Asp Trp Ile Ile Ala Pro Leu Asp Tyr Glu Ala Tyr His Cys Glu Gly

Val Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala

Ile Ile Gln Thr Leu Leu Asn Ser Met Ala Pro Asp Ala Ala Pro Ala 50 55

Ser Cys Cys Val Pro Ala Arg Leu Ser Pro Ile Ser Ile Leu Tyr Ile 75

Asp Ala Ala Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val 85 90

Glu Ala Cys Gly Cys Arg 100

<210> 88

<211> 102

<212> PRT

<213> Homo sapiens

<220>

<223> CDMP-3 construct

<400> 88

Cys Ser Arg Lys Pro Leu His Val Asp Phe Lys Glu Leu Gly Trp Asp 1 10 15

Asp Trp Ile Ile Ala Pro Leu Asp Tyr Glu Ala Tyr His Cys Glu Gly
20 25 30

Leu Cys Asp Phe Pro Leu Arg Ser His Leu Glu Pro Thr Asn His Ala 35 40 45

Ile Ile Gln Thr Leu Leu Asn Ser Met Ala Pro Asp Ala Ala Pro Ala 50 55 60

Ser Cys Cys Val Pro Ala Arg Leu Ser Pro Ile Ser Ile Leu Tyr Ile 65 70 75 80

Asp Ala Ala Asn Asn Val Val Tyr Lys Gln Tyr Glu Asp Met Val Val 85 90 95

Glu Ala Cys Gly Cys Arg 100

<210> 89

<211> 129

<212> PRT

<213> Homo sapiens

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<223> H2487

<400> 89

Met Thr Met Ile Thr Asn Ser Leu Ala Ser Trp Arg Glu Pro Ser Phe 1 5 10 15

Met Ala Leu Ser Ser Ser Asp Gln Arg Gln Ala Cys Lys Lys His Glu 20 25 30

Leu Ty	r Val 35	Ser	Phe	Arg	Asp	Leu 40	Gly	Trp	Gln	Asp	Trp 45	Ile	Ile	Ala	
Pro Gl 50	_	Tyr	Ala	Ala	Tyr 55	туr	Суз	Glu	Gly	Glu 60	Сув	Ala	Phe	Pro	
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Page 61

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